=> file biosis medline caplus wpids uspatfull COST IN U.S. DOLLARS

10/648,740

SINCE FILE

TOTAL SESSION 1.32

ENTRY

0.21

FULL ESTIMATED COST

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FILE 'MEDLINE' ENTERED AT 11:17:52 ON 14 FEB 2006

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FILE 'USPATFULL' ENTERED AT 11:17:52 ON 14 FEB 2006
CA INDEXING COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

*** YOU HAVE NEW MAIL ***

=> s synthes? (3a) oligo?
L1 71976 SYNTHES? (3A) OLIGO?

=> s l1 and solid suppot L2 0 L1 AND SOLID SUPPOT

=> s l1 and solid support L3 19485 L1 AND SOLID SUPPORT

=> s 13 and carbonate (4a) protect? (3a) group?
L4 25 L3 AND CARBONATE (4A) PROTECT? (3A) GROUP?

=> s 14 and simultan?

L5 21 L4 AND SIMULTAN?

=> s 15 and phosphite triester L6 16 L5 AND PHOSPHITE TRIESTER

=> dup rem 16
PROCESSING COMPLETED FOR L6

L7 16 DUP REM L6 (0 DUPLICATES REMOVED)

=> d 17 bib abs 1-16

L7 ANSWER 1 OF 16 USPATFULL on STN

AN 2005:57493 USPATFULL

TI Exocyclic amine triaryl methyl protecting groups in two step

polynucleotide synthesis

IN Dellinger, Douglas J., Boulder, CO, UNITED STATES Sierzchala, Agnieszka B., Boulder, CO, UNITED STATES Caruthers, Marvin H., Boulder, CO, UNITED STATES

PI US 2005049411 A1 20050303

AI US 2003-652064 A1 20030830 (10)

DT Utility

FS APPLICATION

LREP AGILENT TECHNOLOGIES, INC., Legal Department, DL429, Intellectual Property Administration, P.O. Box 7599, Loveland, CO, 80537-0599

CLMN Number of Claims: 21 ECL Exemplary Claim: 1 DRWN 3 Drawing Page(s)

LN.CNT 1531

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Precursors for use in the synthesis of polynucleotides and methods of using the precursors in synthesizing polynucleotides are disclosed. The precursors include a heterocyclic base having an exocyclic amine group

and a substituted or unsubstituted triaryl methyl protecting group bound to the exocyclic amine group.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

```
L7
     ANSWER 2 OF 16 USPATFULL on STN
       2005:57489 USPATFULL
AN
ΤI
       Precursors for two-step polynucleotide synthesis
IN
       Dellinger, Douglas J., Boulder, CO, UNITED STATES
       Sierzchala, Agnieszka B., Boulder, CO, UNITED STATES
       Caruthers, Marvin H., Boulder, CO, UNITED STATES
                         A1
                               20050303
PΙ
       US 2005049407
AΙ
       US 2003-652048
                          A1
                               20030830 (10)
DT
       Utility
       APPLICATION
FS
       AGILENT TECHNOLOGIES, INC., Legal Department, DL429, Intellectual
LREP
       Property Administration, P.O. Box 7599, Loveland, CO, 80537-0599
CLMN
       Number of Claims: 26
ECL
       Exemplary Claim: 1
       3 Drawing Page(s)
DRWN
LN.CNT 1564
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Precursors for use in the synthesis of polynucleotides are disclosed.
       The precursors include a heterocyclic base having an exocyclic amine
       group and a substituted or unsubstituted triaryl methyl protecting group
       bound to the exocyclic amine group.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
L7
     ANSWER 3 OF 16 USPATFULL on STN
AN
       2005:56686 USPATFULL
ΤI
       Method for polynucleotide synthesis
       Dellinger, Douglas J., Boulder, CO, UNITED STATES
IN
       Dellinger, Geraldine, Boulder, CO, UNITED STATES
       Sierzchala, Agnieszka B., Boulder, CO, UNITED STATES
       Caruthers, Marvin H., Boulder, CO, UNITED STATES
       US 2005048601
                         A1
                               20050303
AΤ
       US 2003-652054
                          A1
                               20030830 (10)
DT
       Utility
FS
      APPLICATION
LREP
      AGILENT TECHNOLOGIES, INC., Legal Department, DL429, Intellectual
       Property Administration, P.O. Box 7599, Loveland, CO, 80537-0599
CLMN
      Number of Claims: 35
ECL
       Exemplary Claim: 1
       3 Drawing Page(s)
DRWN
LN.CNT 2443
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
      Methods of forming an internucleotide bond are disclosed. Such methods
       find use in synthesis of polynucleotides. The method involves contacting
       a functionalized support with a precursor having an exocyclic amine
       triaryl methyl protecting group under conditions and for a time
```

sufficient to result in internucleotide bond formation. The

site hydroxyl, the nucleoside moiety attached to the solid

triaryl methyl linker group, and a nucleoside moiety having a reactive

functionalized support includes a solid support, a

support via the triaryl methyl linker group.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L7

```
ANSWER 4 OF 16 USPATFULL on STN
AN
      2005:56582 USPATFULL
ΤI
      Cleavable linker for polynucleotide synthesis
TN
      Dellinger, Douglas J., Boulder, CO, UNITED STATES
      Dellinger, Geraldine, Boulder, CO, UNITED STATES
      Caruthers, Marvin H., Boulder, CO, UNITED STATES
PΙ
      US 2005048497
                      A1
                              20050303
ΑI
      US 2003-652063
                         A1
                              20030830 (10)
ĎΤ
      Utility
FS
      APPLICATION
```

```
Property Administration, P.O. Box 7599, Loveland, CO, 80537-0599
CLMN
       Number of Claims: 23
       Exemplary Claim: 1
ECL
DRWN
       2 Drawing Page(s)
LN.CNT 1803
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Functionalized supports for polynucleotide synthesis are disclosed. The
AB
       supports have linker moieties that are stable to conditions used in
       polynucleotide synthesis, but may be cleaved to release synthesized
       polynucleotides from the support. Methods of making the functionalized
       supports and methods of using are also disclosed.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
     ANSWER 5 OF 16 USPATFULL on STN
L7
       2005:56581 USPATFULL
AN
       Method of polynucleotide synthesis using modified support
ΤI
       Dellinger, Douglas J., Boulder, CO, UNITED STATES
IN
       Dellinger, Geraldine, Boulder, CO, UNITED STATES
       Hargreaves, John, Mountain View, CA, UNITED STATES
       US 2005048496
                          A1
                               20050303
PΙ
       US 2003-652049
                          A1
                               20030830 (10)
ΑI
DT
       Utility
FS
       APPLICATION
LREP
       AGILENT TECHNOLOGIES, INC., Legal Department, DL429, Intellectual
       Property Administration, P.O. Box 7599, Loveland, CO, 80537-0599
CLMN
       Number of Claims: 31
ECL
       Exemplary Claim: 1
DRWN
       3 Drawing Page(s)
LN.CNT 2081
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Methods for polynucleotide synthesis using modified support materials
       are disclosed. The synthesis reaction typically involves concurrent
       oxidation and deprotection reactions. Upon synthesis of a desired
       polynucleotide, the completed polynucleotide may be released from the
       modified support materials.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
     ANSWER 6 OF 16 USPATFULL on STN
1.7
       2004:292960 USPATFULL
AN
       Methods of synthesizing oligonucleotides using
TI
       carbonate protecting groups and alpha-effect
       nucleophile deprotection
       Dellinger, Douglas J., Sunnyvale, CA, UNITED STATES
IN
       Caruthers, Marvin H., Boulder, CO, UNITED STATES
       Betley, Jason R., Edmunds Suffolk, UNITED KINGDOM
PΙ
       US 2004230052
                          A1
                               20041118
ΑI
       US 2003-648740
                          Α1
                               20030825 (10)
RLI
       Continuation of Ser. No. US 2001-756991, filed on 8 Jan 2001, GRANTED,
       Pat. No. US 6630581 Division of Ser. No. US 1999-338179, filed on 22 Jun
       1999, GRANTED, Pat. No. US 6222030 Continuation-in-part of Ser. No. US
       1998-128052, filed on 3 Aug 1998, ABANDONED
DT
       Utility
FS
       APPLICATION
LREP
       AGILENT TECHNOLOGIES, INC., INTELLECTUAL PROPERTY ADMINISTRATION, LEGAL
       DEPT., P.O. BOX 7599, M/S DL429, LOVELAND, CO, 80537-0599
CLMN
       Number of Claims: 30
ECL
       Exemplary Claim: 1
DRWN
       7 Drawing Page(s)
LN.CNT 1411
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AΒ
       The invention provides methods for synthesizing
       oligonucleotides using nucleoside monomers having
       carbonate protected hydroxyl groups that are
       deprotected with \alpha-effect nucleophiles. The \alpha-effect
       nucleophile irreversibly cleave the carbonate
       protecting groups while simultaneously
```

AGILENT TECHNOLOGIES, INC., Legal Department, DL429, Intellectual

LREP

oxidizing the internucleotide phosphite triester linkage to a phosphodiester linkage. The procedure may be carried out in aqueous solution at neutral to mildly basic pH. The method eliminates the need for separate deprotection and oxidation steps, and, since the use of acid to remove protecting groups is unnecessary, acid-induced depurination is avoided. Fluorescent or other readily detectable carbonate protecting groups can be used, enabling monitoring of individual reaction steps during oligonucleotide synthesis. The invention is particularly useful in the highly parallel, microscale synthesis of oligonucleotides.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

```
ANSWER 7 OF 16 USPATFULL on STN
L7
       2003:214617 USPATFULL
AN
      Process for the synthesis of oligomeric compounds
ΤI
      Cheruvallath, Zacharia S., San Diego, CA, UNITED STATES
TN
      Ravikumar, Vasulinga T., Carlsbad, CA, UNITED STATES
      Cole, Douglas L., San Diego, CA, UNITED STATES
       ISIS Pharmaceuticals, Inc., Carlsbad, CA (U.S. corporation)
PA
      US 2003149260
                               20030807
PI
                        A1
      US 6677471
                         B2
                               20040113
      US 2002-290587
                               20021108 (10)
ΑI
                         A1
RLI
      Continuation of Ser. No. US 2001-16465, filed on 11 Dec 2001, GRANTED,
       Pat. No. US 6521775 Division of Ser. No. US 1999-349659, filed on 8 Jul
       1999, GRANTED, Pat. No. US 6399756 Continuation-in-part of Ser. No. US
       1998-111678, filed on 8 Jul 1998, GRANTED, Pat. No. US 6326478
DT
      Utility
      APPLICATION
FS
LREP
      WOODCOCK WASHBURN LLP, ONE LIBERTY PLACE, 46TH FLOOR, 1650 MARKET
      STREET, PHILADELPHIA, PA, 19103
      Number of Claims: 57
CLMN
ECL
      Exemplary Claim: 1
DRWN
      No Drawings
LN.CNT 2248
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
      Synthetic processes are provided wherein oligomeric compounds are
AB
      prepared having phosphodiester, phosphorothioate, phosphorodithioate, or
      other covalent linkages. Also provided are synthetic intermediates
      useful in such processes.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
```

```
ANSWER 8 OF 16 USPATFULL on STN
L7
      2003:38365 USPATFULL
AN
ΤI
      Polynucleotide synthesis
      Perbost, Michel G.M., Cupertino, CA, UNITED STATES
IN
PΙ
      US 2003028012
                       A1
                               20030206
                               20020917 (10)
ΑI
      US 2002-245211
                         A1
      Continuation of Ser. No. US 1999-420099, filed on 18 Oct 1999, GRANTED,
RLI
      Pat. No. US 6451998
DT
      Utility
FS
      APPLICATION
LREP
      AGILENT TECHNOLOGIES, INC., Legal Department, DL429, Intellectual
      Property Administration, P.O. Box 7599, Loveland, CO, 80537-0599
CLMN
      Number of Claims: 24
ECL
      Exemplary Claim: 1
DRWN
      4 Drawing Page(s)
LN.CNT 748
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
      A method including coupling the moiety to a phospho or phosphite
AB
      derivative of a protected alcohol, so as to form the corresponding
```

phosphate or phosphite between the hydroxy and phospho or phosphite groups. The hydroxy group may be later de-protected by hydrolyzing the resulting compound to deprotect the protected alcohol and cleave the phosphate from the moiety so as to regenerate the hydroxy group of the moiety. The method has particular application to fabrication of addressable polynucleotide arrays and allows failed sequences, as well as inter-feature regions, to be left with a free hydroxy group at the ends of the molecules (failed sequences or linkers) at such locations.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

```
L7
     ANSWER 9 OF 16 USPATFULL on STN
AN
       2002:113056 USPATFULL
ΤI
       Synthesis of polynucleotides using combined oxidation/deprotection
       chemistry
ΙN
       Dellinger, Douglas J., Sunnyvale, CA, UNITED STATES
       Perbost, Michael G. M., Bethany, CT, UNITED STATES
       Caruthers, Marvin H., Boulder, CO, UNITED STATES
       Betley, Jason R., Suffolk, UNITED KINGDOM
PΙ
       US 2002058802
                          A1
                               20020516
       US 2001-916369
ΑI
                          A1
                               20010727 (9)
       Continuation-in-part of Ser. No. US 2000-627249, filed on 28 Jul 2000,
RLI
       PENDING
DT
       Utility
FS
       APPLICATION
       AGILENT TECHNOLOGIES, INC., Legal Department, DL429, Intellectual
LREP
       Property Administration, P.O. Box 7599, Loveland, CO, 80537-0599
CLMN
       Number of Claims: 25
ECL
       Exemplary Claim: 1
DRWN
       4 Drawing Page(s)
LN.CNT 1957
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       A method of synthesizing a polynucleotide which can, for example, be
       used during fabrication of an array. A second nucleoside is coupled to a
       first nucleoside through a phosphite linkage, with the second nucleoside
```

having a hydroxyl protecting group that is a non-carbonate protecting group. The product of the foregoing step is

phosphate and deprotects the protected hydroxyl of the coupled nucleoside. The method has particular application to fabricating an addressable array of polynucleotides on a substrate which carries

exposed to a composition which both oxidizes the formed phosphite to a

substrate bound moieties each with a hydroxyl group.

```
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
L7
     ANSWER 10 OF 16 USPATFULL on STN
AN
       2002:106412 USPATFULL
TI
       Process for the synthesis of oligomeric compounds
IN
       Cheruvallath, Zacharia S., San Diego, CA, UNITED STATES
       Ravikumar, Vasulinga T., Carlsbad, CA, UNITED STATES
       Cole, Douglas L., San Diego, CA, UNITED STATES
       ISIS Pharmaceuticals. Inc. (U.S. corporation)
PA
ΡI
       US 2002055623
                       A1
                               20020509
       US 6521775
                         B2
                               20030218
AΤ
       US 2001-16465
                         A1
                               20011211 (10)
RLI
       Division of Ser. No. US 1999-349659, filed on 8 Jul 1999, PENDING
       Continuation-in-part of Ser. No. US 1998-111678, filed on 8 Jul 1998,
       PATENTED
       Utility
DT
FS
       APPLICATION
LREP
       WOODCOCK WASHBURN LLP, One Liberty Place - 46th Floor, Philadelphia, PA,
       19103
       Number of Claims: 57
CLMN
ECL
       Exemplary Claim: 1
DRWN
       No Drawings
LN.CNT 2243
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AB
       Synthetic processes are provided wherein oligomeric compounds are
       prepared having phosphodiester, phosphorothioate, phosphorodithioate, or
       other covalent linkages. Also provided are synthetic intermediates
       useful in such processes.
```

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

```
AN
       2002:85181 USPATFULL
TI
       Solid phase synthesis of oligonucleotides using
       carbonate protecting groups and alpha-effect
       nucleophile deprotection
       Dellinger, Douglas J., Sunnyvale, CA, UNITED STATES
IN
       Caruthers, Marvin H., Boulder, CO, UNITED STATES
       Betley, Jason R., Bury St. Edmonds, UNITED KINGDOM
ΡI
       US 2002045221
                         A1
                               20020418
       US 6630581
                         B2
                               20031007
ΑI
       US 2001-756991
                         A1
                               20010108 (9)
       Division of Ser. No. US 1999-338179, filed on 22 Jun 1999, UNKNOWN
RLI
DT
       Utility
FS
       APPLICATION
       AGILENT TECHNOLOGIES, Legal Department, 51 U-PD, Intellectual Property
LREP
       Administration, P. O. Box 58043, Santa Clara, CA, 95052-8043
       Number of Claims: 52
CLMN
       Exemplary Claim: 1
ECL
       7 Drawing Page(s)
DRWN
LN.CNT 1526
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention provides a method for synthesizing
       oligonucleotides using carbonate protection
       of hydroxyl groups and nucleophilic deprotection reagents. The
       deprotection reagents irreversibly cleave the carbonate
       protecting groups while simultaneously
       oxidizing the internucleotide phosphite triester
       linkage, and can be used in aqueous solution at neutral to mildly basic
       pH. The method eliminates the need for separate deprotection and
       oxidation steps, and, since the use of acid to remove protecting groups
       is unnecessary, acid-induced depurination is avoided. Fluorescent or
       other readily detectable carbonate protecting
       groups can be used, enabling monitoring of individual reaction
       steps during oligonucleotide synthesis. The
       invention is particularly useful in the highly parallel, microscale
       synthesis of oligonucleotides. Reagents and kits for
       carrying out the aforementioned method are provided as well.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
L7
     ANSWER 12 OF 16 USPATFULL on STN
AN
       2002:239168 USPATFULL
TI
       Capping and de-capping during oligonucleotide
       synthesis
IN
       Perbost, Michael G. M., Cupertino, CA, United States
PA
       Agilent Technologies, Inc., Palo Alto, CA, United States (U.S.
       corporation)
                               20020917
PΙ
       US 6451998
                          В1
ΑI
       US 1999-420099
                               19991018 (9)
DT
       Utility
       GRANTED
FS
EXNAM Primary Examiner: Richter, Johann; Assistant Examiner: Crane, L. Eric
LREP
       Stewart, Gordon M.
CLMN
       Number of Claims: 24
ECL
       Exemplary Claim: 10,11
DRWN
       7 Drawing Figure(s); 4 Drawing Page(s)
LN.CNT 770
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AB
       A method of capping a hydroxy group of a moiety, comprising coupling the
       moiety to a phosphor or phosphite derivative of a protected alcohol, so
       as to form the corresponding phosphate or phosphite between the hydroxy
       and phosphor or phosphite groups. The hydroxy group may be later
       de-capped by hydrolyzing the resulting compound to deprotect the
       protected alcohol and cleave the phosphate from the moiety so as to
       regenerate the hydroxy group of the moiety. The method has particular
       application to fabrication of addressable polynucleotide arrays and
       allows failed sequences, as well as inter-feature regions, to be left
       with a free hydroxy group at the ends of the molecules (failed sequences
```

or linkers) at such locations.

```
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
L7
     ANSWER 13 OF 16 USPATFULL on STN
       2002:130084 USPATFULL
AN
       Process for the synthesis of oligomeric compounds
ТT
IN
       Cheruvallath, Zacharia S., San Diego, CA, United States
       Ravikumar, Vasulinga T., Carlsbad, CA, United States
       Cole, Douglas L., San Diego, CA, United States
       ISIS Pharmaceuticals, Inc., Carlsbad, CA, United States (U.S.
PA
       corporation)
PΙ
       US 6399756
                          B1
                               20020604
       US 1999-349659
                               19990708 (9)
AΙ
RLI
       Continuation-in-part of Ser. No. US 1998-111678, filed on 8 Jul 1998,
       now abandoned
DT
       Utility
FS
       GRANTED
       Primary Examiner: Geist, Gary; Assistant Examiner: Crane, L. E.
EXNAM
       Woodcock Washburn LLP
LREP
CLMN
       Number of Claims: 52
ECL
       Exemplary Claim: 1
DRWN
       0 Drawing Figure(s); 0 Drawing Page(s)
LN.CNT 2423
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AB
       Synthetic processes are provided wherein oligomeric compounds are
       prepared having phosphodiester, phosphorothioate, phosphorodithioate, or
       other covalent linkages. Also provided are synthetic intermediates
       useful in such processes.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
L7
    ANSWER 14 OF 16 USPATFULL on STN
AN
       2001:60049 USPATFULL
       Solid phase synthesis of oligonucleotides using
TΙ
       carbonate protecting groups and alpha-effect
       nucleophile deprotection
IN
       Dellinger, Douglas J., Sunnyvale, CA, United States
       Caruthers, Marvin H., Boulder, CO, United States
       Betley, Jason R., Bury St. Edmunds, United Kingdom
       Agilent Technologies, Inc., Palo Alto, CA, United States (U.S.
PA
       corporation)
PΙ
       US 6222030
                          B1
                               20010424
                               19990622 (9)
       US 1999-338179
AΤ
RLI
       Continuation-in-part of Ser. No. US 1998-128052, filed on 3 Aug 1998
DT
       Utility
FS
       Granted
EXNAM Primary Examiner: Riley, Jezia
       Number of Claims: 39
CLMN
ECL
       Exemplary Claim: 1
DRWN
       8 Drawing Figure(s); 7 Drawing Page(s)
LN.CNT 1476
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention provides a method for synthesizing
AB
       oligonucleotides using carbonate protection
       of hydroxyl groups and nucleophilic deprotection reagents. The
       deprotection reagents irreversibly cleave the carbonate
       protecting groups while simultaneously
       oxidizing the intemucleotide phosphite triester
       linkage, and can be used in aqueous solution at neutral to mildly basic
       pH. The method eliminates the need for separate deprotection and
       oxidation steps, and, since the use of acid to remove protecting groups
       is unnecessary, acid-induced depurination is avoided. Fluorescent or
       other readily detectable carbonate protecting
       groups can be used, enabling monitoring of individual reaction
       steps during oligonucleotide synthesis. The
       invention is particularly useful in the highly parallel, microscale
```

synthesis of oligonucleotides. Reagents and kits for

carrying out the aforementioned method are provided as well.

```
*ANSWER 15 OF 16 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN
     2000-225901 [20]
                        WPIDS
AN
CR
     2005-513718 [53]
DNC C2000-069092
     Oligonucleotide synthesis by phosphoramidite method
TI
     using carbonate protecting group and
     deprotecting reagent that simultaneously oxidizes
     phosphite triester linkage to phosphotriester linkage.
חכ
     B04 D16 J04
IN
     BETLEY, J R; CARUTHERS, M H; DELLINGER, D J
     (AGIL-N) AGILENT TECHNOLOGIES INC; (HEWP) HEWLETT-PACKARD CO; (BETL-I)
PA
     BETLEY J R; (CARU-I) CARUTHERS M H; (DELL-I) DELLINGER D J
CYC
     26
                     A2 20000308 (200020)* EN
PΙ
     EP 984021
         R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT
            RO SE SI
     US 6222030
                    B1 20010424 (200125)
     US 2002045221 A1 20020418 (200228)
                    B2 20031007 (200374)
     US 6630581
     US 2004230052 A1 20041118 (200477)
                    B1 20050427 (200532)
     EP 984021
         R: DE FR GB
     DE 69924930
                    E 20050602 (200538)
     EP 984021 A2 EP 1999-306168 19990803; US 6222030 B1 CIP of US 1998-128052
ADT
     19980803, US 1999-338179 19990622; US 2002045221 A1 Div ex US 1999-338179
     19990622, US 2001-756991 20010108; US 6630581 B2 CIP of US 1998-128052
     19980803, Div ex US 1999-338179 19990622, US 2001-756991 20010108; US
     2004230052 A1 CIP of US 1998-128052 19980803, Div ex US 1999-338179
     19990622, Cont of US 2001-756991 20010108, US 2003-648740 20030825; EP
     984021 B1 EP 1999-306168 19990803; DE 69924930 E DE 1999-624930 19990803,
     EP 1999-306168 19990803
     US 6630581 B2 Div ex US 6222030; US 2004230052 A1 Div ex US 6222030, Cont
     of US 6630581; DE 69924930 E Based on EP 984021
PRAI US 1999-338179
                          19990622; US 1998-128052
                                                         19980803;
     US 2001-756991
                          20010108; US 2003-648740
                                                         20030825
     2000-225901 [20]
                        WPIDS
AN
CR
     2005-513718 [53]
           984021 A UPAB: 20050818
AΒ
     NOVELTY - Oligonucleotide synthesis comprises
     condensing the 3'- or 5'-OH group of a supported (oligo) nucleoside with a
     nucleoside phosphoramidite having a protected OH group, to form an
     intermediate where the (oligo) nucleoside is bound to the nucleoside by a
     phosphite triester linkage; and deprotecting the
     intermediate with a reagent which also oxidizes the phosphite
     triester linkage.
          DETAILED DESCRIPTION - Oligonucleotide synthesis
     process comprises:
          (a) condensing the 3'- or 5'-hydroxy group of a support-bound
     nucleoside or oligonucleotide with a monomeric nucleoside phosphoramidite
     having a carbonate-protected hydroxy group,
     to form an intermediate in which the support-bound nucleoside or
     oligonucleotide is bound to the monomeric nucleoside through a
     phosphite triester linkage; and
          (b) treating the intermediate with a deprotecting reagent capable of
     removing the carbonate protecting group and
     simultaneously oxidizing the phosphite triester
     linkage to a phosphotriester linkage.
          INDEPENDENT CLAIMS are also included for the following:
          (A) a method for making an oligonucleotide array made up of array
     features, each presenting a specified oligonucleotide sequence at an
     address on a substrate, comprising: providing a hydroxyl-derivatized array
     substrate and treating the array substrate to protect the hydroxyl groups
     on the derivatized surface from reaction with phosphoramidites; and then
     iteratively carrying out the steps of (i) applying droplets of an alpha
     effect nucleophile to effect deprotection of the hydroxyl groups at
     selected addresses, and (ii) flooding the array substrate with a medium
     containing a selected monomeric nucleoside phosphoramidite having a
```

carbonate-protected hydroxyl group, to permit

covalent attachment of the selected nucleoside to the deprotected hydroxyl groups at the selected addresses;

(B) a kit for synthesizing an oligonucleotide on a solid support, comprising: a hydroxyl-derivatized support surface; a protecting group for protecting hydroxyl groups on the derivatized support surface; at least one protected nucleoside; at least one nucleoside phosphoramidite; a nucleophile that exhibits an alpha effect at neutral to mildly basic pH; and reagents suitable for establishing pH and for carrying out reactions of deprotection, phosphoramidite coupling and oxidation to form an internucleotide phosphotriester linkage; and

(C) a nucleoside monomer of formula (I) or (II).

B = a purine or pyrimidine base;

R = H or OH;

(C) a nucleoside monomer of formula (1) or (11)

B = a purine or pyrimidine base;

R = H or OH;

R1 = COOR3;

R3 = optionally substituted hydrocarbyl;

R2 = a group of formula (i);

X = NQ1Q2;

Q1, Q2 = alkyl, aryl, aralkyl, alkaryl, cycloalkyl, alkenyl, cycloalkenyl, alkynyl or cycloalkynyl, optionally containing one or more nonhydrocarbyl linkages and optionally C-substituted by nonhydrocarbyl substituents, or are linked to form a mono- or polyheterocyclic ring; and Y = H or hydrocarbyl.

USE - The process has utility in the fields of biochemistry, molecular biology, pharmacology, and medical diagnostics and screening. ADVANTAGE - The carbonate protecting

group can be removed at neutral to mildly basic pH, thus avoiding acid-induced depurination. Deprotection and phosphite oxidation are effected simultaneously, eliminating the need for a separate oxidation step. The deprotecting agent also removes exocyclic amine protecting groups. Detectable protecting groups can be used to enable monitoring of individual reaction steps. The process can be used to perform 3'-to-5' or 5'-to-3' syntheses and readily lends itself to highly parallel microscale oligonucleotide synthesis.

Dwg.0/7

```
L7
     ANSWER 16 OF 16 USPATFULL on STN
ΑN
       1999:63399 USPATFULL
       5'to 3' nucleic acid synthesis using 3'-photoremovable protecting group
TI
IN
       Pirrung, Michael C., Houston, TX, United States
       Shuey, Steven W., Durham, NC, United States
       Bradley, Jean-Claude, Durham, NC, United States
PA
       Duke University, Durham, NC, United States (U.S. corporation)
PΙ
       US 5908926
                               19990601
ΑI
      US 1995-406327
                               19950316 (8)
      Utility
DT
FS
       Granted
EXNAM Primary Examiner: Kunz, Gary L.
LREP
      Nixon & Vanderhye P.C.
CLMN
      Number of Claims: 20
ECL
      Exemplary Claim: 1
DRWN
      No Drawings
LN.CNT 635
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
```

The present invention relates, in general, to a method of synthesizing a nucleic acid, and, in particular, to a method of effecting 5' to 3' nucleic acid synthesis. The method can be used to prepare arrays of oligomers bound to a support via their 5' end. The invention also relates to a method of effecting mutation analysis using such arrays.

The invention further relates to compounds and compositions suitable for use in such methods.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L1 L2

L3

L4 L_5

L6 L7

L8

L9

AN

TΙ

IN

PΑ

PΤ

AΙ

DT

FS

AB

ΑN

DN

TI

```
(FILE 'HOME' ENTERED AT 11:14:26 ON 14 FEB 2006)
     FILE 'STNGUIDE' ENTERED AT 11:17:30 ON 14 FEB 2006
     FILE 'HOME' ENTERED AT 11:17:35 ON 14 FEB 2006
     FILE 'BIOSIS, MEDLINE, CAPLUS, WPIDS, USPATFULL' ENTERED AT 11:17:52 ON
     14 FEB 2006
          71976 S SYNTHES? (3A) OLIGO?
              0 S L1 AND SOLID SUPPOT
          19485 S L1 AND SOLID SUPPORT
             25 S L3 AND CARBONATE (4A) PROTECT? (3A) GROUP?
             21 S L4 AND SIMULTAN?
             16 S L5 AND PHOSPHITE TRIESTER
             16 DUP REM L6 (0 DUPLICATES REMOVED)
              9 S L4 NOT L7
              7 DUP REM L8 (2 DUPLICATES REMOVED)
=> s 14 and phosphite
            21 L4 AND PHOSPHITE
=> s 110 not 17
             5 L10 NOT L7
L11
=> dup rem 111
PROCESSING COMPLETED FOR L11
              3 DUP REM L11 (2 DUPLICATES REMOVED)
1.12
=> d l12 bib abs 1-3
L12 ANSWER 1 OF 3 USPATFULL on STN
       2004:314490 USPATFULL
       Releasable polymer arrays
       Cuppoletti, Andrea, Livermore, CA, UNITED STATES
       McGall, Glenn H., Palo Alto, CA, UNITED STATES
       Affymetrix, INC., Santa Clara, CA (U.S. corporation)
                               20041209
       US 2004248162
                         A1
       US 2004-791005
                               20040302 (10)
                         A1
       Continuation-in-part of Ser. No. US 2003-738381, filed on 16 Dec 2003,
RLI
       PENDING
      US 2002-434144P
PRAI
                          20021217 (60)
       Utility
       APPLICATION
       AFFYMETRIX, INC, ATTN: CHIEF IP COUNSEL, LEGAL DEPT., 3380 CENTRAL
LREP
       EXPRESSWAY, SANTA CLARA, CA, 95051
CLMN
       Number of Claims: 32
ECL
       Exemplary Claim: 1
DRWN
       No Drawings
LN.CNT 1394
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Methods are provided for fabricating an array of polymers wherein the
       polymers may be released from the surface of the array by activation of
       a cleavable moiety. Also provided are arrays of polymers having of
       polymers wherein the polymers can be released from the surface of the
       array by activation of a releasable group. Arrays of nucleic acids
       wherein a nucleic acid probe may be released from the array by
       activation of a releasable groups and methods for fabrication of such
       arrays are also disclosed.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
L12 ANSWER 2 OF 3 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN
     DUPLICATE 1
     2004:144635 BIOSIS
     PREV200400146581
```

Solid-phase oligodeoxynucleotide synthesis: A two-step

cycle using peroxy anion deprotection. ΑU *Sierzchala, Agnieszka B.; Dellinger, Douglas J.; Betley, Jason R.; Wyrzykiewicz, Tadeusz K.; Yamada, Christina M.; Caruthers, Marvin H. [Reprint Author] Department of Chemistry and Biochemistry, University of Colorado, Boulder, CS CO, 80309, USA doug dellinger@agilent.com; marvin.caruthers@colorado.edu Journal of the American Chemical Society, (November 5 2003) Vol. 125, No. SO 44, pp. 13427-13441. print. ISSN: 0002-7863 (ISSN print). Article DT English ED Entered STN: 17 Mar 2004

Last Updated on STN: 17 Mar 2004 AΒ A novel solid-phase phosphoramidite based oligodeoxynucleotide two-step synthesis method has been developed. Keys to this method are replacement of the 5'-dimethoxytrityl blocking group with an aryloxycarbonyl and the use of N-dimethoxytrityl protection for the exocyclic amines of adenine and cytosine. With these modifications, coupling of each 2'-deoxynucleoside 3'-phosphoramidite to the growing oligodeoxynucleotide on the solid support can be followed by treatment with an aqueous mixture of peroxy anions buffered at This reagent effectively removes the carbonate protecting group and simultaneously oxidizes the phosphite internucleotide linkage. As a consequence a new two-step synthesis cycle is possible. Oligodeoxynucleotides synthesized using this approach are identical to authentic samples when tested by a variety of analytical techniques.

L12 ANSWER 3 OF 3 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN 2002-156732 [21] ANCR 2002-499520 [53]

DNC C2002-049016

Synthesis of polynucleotide useful during fabrication of an array involves coupling nucleoside phosphoramidite and a solid-supported nucleoside and treating the product with an oxidation/deprotection composition.

DC B04 D16

IN BETLEY, J R; CARUTHERS, M; DELLINGER, D J; PERBOST, M G M

PA (AGIL-N) AGILENT TECHNOLOGIES INC

CYC 26

PТ EP 1176151 A1 20020130 (200221)* EN 36

> R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI TR

ADT EP 1176151 A1 EP 2001-118360 20010727

PRAI US 2000-627249 20000728

2002-156732 [21] WPIDS $\mathbf{A}\mathbf{N}$

CR 2002-499520 [53]

AΒ 1176151 A UPAB: 20020823

> NOVELTY - Synthesis of a polynucleotide involves coupling a second nucleoside to a first nucleoside through a phosphite linkage, where the second nucleoside has a non-carbonate protecting group protecting a hydroxyl, and exposing the product to a composition which concurrently oxidizes the phosphate formed to a phosphate and deprotects the protected hydroxyl of the second nucleoside.

> DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) a method of fabricating an addressable array of polynucleotides on a substrate carrying substrate bound groups, each with a hydroxyl group, comprising, at each multiple different substrate addresses:
- (a) coupling a second nucleoside to a first nucleoside through a phosphite linkage, where the second nucleoside has a noncarbonate protecting group protecting
- a hydroxyl, and exposing the product to a composition which concurrently oxidizes the phosphate formed to a phosphate and deprotects the protected hydroxyl of the second nucleoside; and
- (b) repeating step (a) where the deprotected hydroxyl of the coupled nucleoside in one cycle serves as the hydroxyl group of substrate bound groups in the next cycle, so as to for the addressable array with

different polynucleotide sequences at different addresses.

- (2) a method for making an oligonucleotide array comprising:
- (a) treating a hydroxyl-derivatized array substrate to protect hydroxyl groups on the derivatized surface from reaction with phosphoramidites;
- (b) applying droplets of an alpha effect nucleophile to carry out deprotection of hydroxyl moieties at the selected addresses; and
- (c) flooding the array substrate with a medium containing a selected monomeric nucleoside phosphoramidite having a carbonate-protected hydroxyl group, to permit covalent attachment of the selected nucleoside to the deprotected hydroxyl groups at the selected addresses.

The oligonucleotide array has array features each presenting a specified oligonucleotide sequence at an address on (c).

USE - The method is useful for synthesizing the polynucleotide; for carrying out either 3' to 5' or 5' to 3' synthesis; and for fabricating an addressable array of polynucleotide on a substrate (claimed).

ADVANTAGE - The method provides concurrent oxidation of the internucleoside linkage and removal of the hydroxyl protecting group, eliminating the extra step present in conventional process for **synthesizing oligonucleotides**. The process requires no washings and the water may optionally be eliminated, the thorough washing to remove water prior to the coupling step in the next cycle is not required or may be reduced. Dwg.0/6